

TREATMENT SET TS332256 - *quercus alba* pollen, *ulmus americana* pollen, *cynodon dactylon* pollen, *phleum pratense* pollen, *acer negundo* pollen, *juniperus ashei* pollen, *populus deltoides* pollen, *fraxinus americana* pollen, *ambrosia artemisiifolia* pollen, *salsola kali* pollen, *rumex acetosella* pollen, *urtica dioica* pollen, *juglans nigra* pollen, *platanus occidentalis* pollen, *carya illinoiensis* pollen, *carya alba* pollen, *amaranthus retroflexus* pollen and *iva annua* var. *annua* pollen injection, solution

TREATMENT SET TS331153 - *chenopodium album* pollen, *amaranthus retroflexus* pollen, *kochia scoparia* pollen, *rumex acetosella* pollen, *salsola kali* pollen, *plantago lanceolata* pollen, *carya illinoiensis* pollen, *carya alba* pollen, *acer negundo* pollen, *fraxinus americana* pollen, *quercus alba* pollen, *betula nigra* pollen, *poa pratensis* pollen, *cynodon dactylon* pollen and *sorghum halepense* pollen injection, solution

TREATMENT SET TS329151 - *chenopodium album* pollen, *amaranthus retroflexus* pollen, *kochia scoparia* pollen, *iva annua* var. *annua* pollen, *ambrosia trifida* pollen, *ambrosia artemisiifolia* pollen, *ambrosia psilostachya* pollen, *rumex acetosella* pollen, *salsola kali* pollen, *plantago lanceolata* pollen, *carya illinoiensis* pollen, *carya alba* pollen, *juniperus virginiana* pollen, *acer negundo* pollen, *fraxinus americana* pollen, *ulmus americana* pollen, *quercus alba* pollen, *populus deltoides* pollen, *betula nigra* pollen, *poa pratensis* pollen, *cynodon dactylon* pollen and *sorghum halepense* pollen injection, solution

TREATMENT SET TS329152 - *fusarium oxysporum*, *cladosporium cladosporioides*, *cochliobolus sativus*, *candida albicans*, *aspergillus fumigatus*, *ustilago cynodontis*, *sporisorium cruentum*, *pleospora herbarum*, *aureobasidium pullulans* var. *pullutans*, *penicillium chrysogenum* var. *chrysogenum*, *periplaneta americana*, *ustilago nuda hordei*, *ustilago maydis*, *ustilago avenae*, *ustilago tritici*, *rhizopus stolonifer*, *phoma destructiva*, *dermatophagoides pteronyssinus*, *canis lupus familiaris* hair, *alternaria alternata*, *dermatophagoides farinae*, *felis catus* hair, *gallus gallus* feather, *anas platyrhynchos* feather and *anser anser* feather injection, solution

TREATMENT SET TS332147 - *amaranthus retroflexus* pollen, *kochia scoparia* pollen, *ambrosia trifida* pollen, *ambrosia artemisiifolia* pollen, *ambrosia psilostachya* pollen, *salsola kali* pollen, *carya illinoiensis* pollen, *carya alba* pollen, *poa pratensis* pollen, *cynodon dactylon* pollen, *candida albicans*, *aspergillus fumigatus*, *sorghum halepense* pollen, *felis catus* hair, *gallus gallus* feather, *anas platyrhynchos* feather and *anser anser* feather injection, solution

TREATMENT SET TS330836 - *salsola kali* pollen, *fraxinus americana* pollen, *quercus alba* pollen, *populus deltoides* pollen and *poa pratensis* pollen injection, solution

TREATMENT SET TS331634 - *dermatophagoides farinae*, *cynodon dactylon* pollen, *alternaria alternata*, *sorghum halepense* pollen, *iva annua* var. *annua* pollen, *cochliobolus sativus*, *phoma destructiva* and *pleospora herbarum* injection, solution

TREATMENT SET TS330927 - *dermatophagoides farinae*, *alternaria alternata*, *canis lupus familiaris* hair, *periplaneta americana*, *gallus gallus* feather, *anas platyrhynchos* feather, *anser anser* feather, *dermatophagoides pteronyssinus*, *candida albicans*, *ustilago nuda hordei*, *ustilago maydis*, *ustilago avenae*, *ustilago tritici*, *pleospora herbarum*, *ustilago cynodontis* and *sporisorium cruentum* injection, solution

TREATMENT SET TS330928 - *ambrosia trifida* pollen, *ambrosia artemisiifolia* pollen, *ambrosia psilostachya* pollen, *cynodon dactylon* pollen, *plantago lanceolata* pollen, *juniperus virginiana* pollen, *acer negundo* pollen, *populus deltoides* pollen, *carya illinoiensis* pollen, *carya alba* pollen, *betula nigra* pollen, *fraxinus americana* pollen, *quercus alba* pollen, *sorghum halepense* pollen, *kochia scoparia* pollen, *chenopodium album* pollen, *amaranthus retroflexus* pollen, *salsola kali* pollen and *rumex acetosella* pollen injection, solution

TREATMENT SET TS331455 - *periplaneta americana*, *gallus gallus* feather, *anas platyrhynchos* feather, *anser anser* feather, *dermatophagoides farinae*, *dermatophagoides pteronyssinus*, *felis catus* hair, *canis lupus familiaris* hair, *betula nigra* pollen, *acer negundo* pollen, *populus deltoides* pollen, *quercus alba* pollen, *carya illinoiensis* pollen, *carya alba* pollen, *ulmus americana* pollen, *ambrosia trifida* pollen, *ambrosia artemisiifolia* pollen, *ambrosia psilostachya* pollen, *chenopodium album* pollen, *rumex acetosella* pollen, *salsola kali* pollen, *plantago lanceolata* pollen, *cynodon dactylon* pollen, *sorghum halepense* pollen, *poa pratensis* pollen, *alternaria alternata*, *aspergillus fumigatus*, *ustilago nuda hordei*, *ustilago maydis*, *ustilago avenae* and *ustilago tritici* injection, solution

TREATMENT SET TS331621 - *felis catus* hair, *poa pratensis* pollen, *periplaneta americana*, *cynodon dactylon* pollen, *dermatophagoides pteronyssinus*, *juniperus virginiana* pollen, *cochliobolus sativus*, *cladosporium cladosporioides*, *penicillium chrysogenum* var. *chrysogenum*, *aureobasidium pullulans* var. *pullutans*, *rhizopus stolonifer*, *acer negundo* pollen, *populus deltoides* pollen, *carya illinoiensis* pollen, *carya alba* pollen, *betula nigra* pollen, *fraxinus americana* pollen, *quercus alba* pollen, *ustilago nuda hordei*, *ustilago maydis*, *ustilago avenae*, *ustilago tritici*, *pleospora herbarum*, *sorghum halepense* pollen, *kochia scoparia* pollen, *chenopodium album* pollen and *iva annua* var. *annua* pollen injection, solution

TREATMENT SET TS330171 - *cynodon dactylon* pollen, *ambrosia artemisiifolia* pollen, *ulmus americana* pollen, *plantago lanceolata* pollen, *sorghum halepense* pollen, *poa pratensis* pollen, *chenopodium album* pollen, *acer negundo* pollen, *festuca pratensis* pollen, *lolium perenne* pollen, *phleum pratense* pollen and *salsola kali* pollen injection, solution

TREATMENT SET TS329197 - *ulmus americana* pollen, *ambrosia trifida* pollen, *ambrosia artemisiifolia* pollen, *ambrosia psilostachya* pollen, *cynodon dactylon* pollen, *poa pratensis* pollen, *betula nigra* pollen, *acer negundo* pollen, *populus deltoides* pollen, *quercus alba* pollen, *carya illinoiensis* pollen, *carya alba* pollen, *sorghum halepense* pollen, *amaranthus retroflexus* pollen, *salsola kali* pollen, *kochia scoparia* pollen and *chenopodium album* pollen injection, solution

TREATMENT SET TS330416 - *ambrosia trifida* pollen, *ambrosia artemisiifolia* pollen, *ambrosia psilostachya* pollen, *canis lupus familiaris* hair, *periplaneta americana*, *gallus gallus* feather, *anas platyrhynchos* feather, *anser anser* feather, *cynodon dactylon* pollen, *plantago lanceolata* pollen, *candida albicans*, *cladosporium cladosporioides*, *penicillium chrysogenum* var.

chrysogenum, phoma destructiva, aureobasidium pullulans var. pullutans, acer negundo pollen, populus deltoides pollen, carya illinoensis pollen, carya alba pollen, betula nigra pollen, quercus alba pollen, pleospora herbarum, sorghum halepense pollen, chenopodium album pollen, iva annua var. annua pollen, amaranthus retroflexus pollen, salsola kali pollen, rumex acetosella pollen, ustilago cynodontis and sporisorium cruentum injection, solution

TREATMENT SET TS329467 - aspergillus fumigatus, fusarium oxysporum, cladosporium cladosporioides, cochliobolus sativus, penicillium chrysogenum var. chrysogenum, phoma destructiva, aureobasidium pullulans var. pullutans, rhizopus stolonifer, acer negundo pollen, carya illinoensis pollen, carya alba pollen, betula nigra pollen, ustilago nuda hordei, ustilago maydis, ustilago avenae, ustilago tritici, pleospora herbarum, kochia scoparia pollen, chenopodium album pollen, iva annua var. annua pollen, rumex acetosella pollen, ustilago cynodontis and sporisorium cruentum injection, solution

TREATMENT SET TS332343 - poa pratensis pollen, ulmus americana pollen, ambrosia trifida pollen, ambrosia artemisiifolia pollen, ambrosia psilostachya pollen, cynodon dactylon pollen, plantago lanceolata pollen, juniperus virginiana pollen, acer negundo pollen, populus deltoides pollen, carya illinoensis pollen, carya alba pollen, betula nigra pollen, fraxinus americana pollen, quercus alba pollen, sorghum halepense pollen, kochia scoparia pollen, chenopodium album pollen, iva annua var. annua pollen, amaranthus retroflexus pollen, salsola kali pollen and rumex acetosella pollen injection, solution

TREATMENT SET TS332344 - felis catus hair, dermatophagoides farinae, alternaria alternata, canis lupus familiaris hair, dermatophagoides pteronyssinus, aspergillus fumigatus, penicillium chrysogenum var. chrysogenum, ustilago nuda hordei, ustilago maydis, ustilago avenae, ustilago tritici, pleospora herbarum, ustilago cynodontis and sporisorium cruentum injection, solution

TREATMENT SET TS329066 - poa pratensis pollen, ambrosia trifida pollen, ambrosia artemisiifolia pollen, ambrosia psilostachya pollen, canis lupus familiaris hair, gallus gallus feather, anas platyrhynchos feather, anser anser feather, dermatophagoides pteronyssinus, plantago lanceolata pollen, candida albicans, juniperus virginiana pollen, aspergillus fumigatus, aureobasidium pullulans var. pullutans, rhizopus stolonifer, acer negundo pollen, fraxinus americana pollen, quercus alba pollen and pleospora herbarum injection, solution

TREATMENT SET TS329915 - felis catus hair, dermatophagoides farinae, alternaria alternata, dermatophagoides pteronyssinus, plantago lanceolata pollen, candida albicans, cladosporium cladosporioides, acer negundo pollen, populus deltoides pollen, betula nigra pollen, sorghum halepense pollen, kochia scoparia pollen and fusarium oxysporum injection, solution

TREATMENT SET TS329109 - alternaria alternata, poa pratensis pollen, lolium perenne pollen, phleum pratense pollen, ulmus americana pollen, acer negundo pollen, quercus alba pollen and ambrosia artemisiifolia pollen injection, solution

TREATMENT SET TS329161 - felis catus hair, dermatophagoides farinae, alternaria alternata, canis lupus familiaris hair, periplaneta americana, dermatophagoides pteronyssinus, candida albicans, aspergillus fumigatus, fusarium oxysporum, cochliobolus sativus, cladosporium cladosporioides, penicillium chrysogenum var. chrysogenum, phoma destructiva, aureobasidium pullulans var. pullutans, rhizopus stolonifer, ustilago nuda hordei, ustilago maydis, ustilago avenae, ustilago tritici, pleospora herbarum, ustilago cynodontis and sporisorium cruentum injection, solution

TREATMENT SET TS329162 - poa pratensis pollen, ambrosia trifida pollen, ambrosia artemisiifolia pollen, ambrosia psilostachya pollen, cynodon dactylon pollen, plantago lanceolata pollen, juniperus virginiana pollen, acer negundo pollen, populus deltoides pollen, carya illinoensis pollen, carya alba pollen, betula nigra pollen, fraxinus americana pollen, quercus alba pollen, sorghum halepense pollen, chenopodium album pollen, iva annua var. annua pollen and amaranthus retroflexus pollen injection, solution

TREATMENT SET TS331975 - poa pratensis pollen, dermatophagoides farinae, ulmus americana pollen, ambrosia trifida pollen, ambrosia artemisiifolia pollen, ambrosia psilostachya pollen, canis lupus familiaris hair, periplaneta americana, dermatophagoides pteronyssinus, plantago lanceolata pollen, candida albicans, juniperus virginiana pollen, acer negundo pollen, populus deltoides pollen, carya illinoensis pollen, carya alba pollen, betula nigra pollen, fraxinus americana pollen, quercus alba pollen, ustilago nuda hordei, ustilago maydis, ustilago avenae, ustilago tritici, chenopodium album pollen, iva annua var. annua pollen, salsola kali pollen and rumex acetosella pollen injection, solution

TREATMENT SET TS329526 - alternaria alternata, ambrosia artemisiifolia pollen, quercus alba pollen, phleum pratense pollen and dermatophagoides farinae injection, solution

TREATMENT SET TS330503 - felis catus hair, poa pratensis pollen, dermatophagoides farinae, alternaria alternata, ulmus americana pollen, ambrosia trifida pollen, ambrosia artemisiifolia pollen, ambrosia psilostachya pollen, canis lupus familiaris hair, periplaneta americana, gallus gallus feather, anas platyrhynchos feather, anser anser feather, cynodon dactylon pollen, juniperus virginiana pollen, aspergillus fumigatus, fusarium oxysporum, cochliobolus sativus, cladosporium cladosporioides, penicillium chrysogenum var. chrysogenum, phoma destructiva, aureobasidium pullulans var. pullutans, pleospora herbarum, sorghum halepense pollen, ustilago cynodontis and sporisorium cruentum injection, solution

TREATMENT SET TS329750 - felis catus hair, poa pratensis pollen, ambrosia trifida pollen, ambrosia artemisiifolia pollen, ambrosia psilostachya pollen, canis lupus familiaris hair, periplaneta americana, cynodon dactylon pollen, dermatophagoides pteronyssinus, juniperus virginiana pollen, cladosporium cladosporioides, acer negundo pollen, populus deltoides pollen, carya illinoensis pollen, carya alba pollen, betula nigra pollen, fraxinus americana pollen, quercus alba pollen, ustilago nuda hordei, ustilago maydis, ustilago avenae, ustilago tritici, pleospora herbarum, sorghum halepense pollen, iva annua var.

*annua* pollen, *amaranthus retroflexus* pollen, *salsola kali* pollen, *rumex acetosella* pollen, *ustilago cynodontis* and *sporisorium cruentum* injection, solution

TREATMENT SET TS331150 - *felis catus* hair, *dermatophagoides farinae*, *ambrosia trifida* pollen, *ambrosia artemisiifolia* pollen, *ambrosia psilostachya* pollen, *canis lupus familiaris* hair, *cynodon dactylon* pollen, *dermatophagoides pteronyssinus*, *juniperus virginiana* pollen, *cochliobolus sativus*, *cladosporium cladosporioides*, *penicillium chrysogenum* var. *chrysogenum*, *phoma destructiva*, *aureobasidium pullulans* var. *pullutans*, *rhizopus stolonifer*, *acer negundo* pollen, *populus deltoides* pollen, *fraxinus americana* pollen, *chenopodium album* pollen, *iva annua* var. *annua* pollen, *amaranthus retroflexus* pollen, *salsola kali* pollen, *ustilago cynodontis* and *sporisorium cruentum* injection, solution

TREATMENT SET TS333176 - *dermatophagoides farinae*, *alternaria alternata*, *canis lupus familiaris* hair, *periplaneta americana*, *dermatophagoides pteronyssinus*, *candida albicans*, *aspergillus fumigatus*, *fusarium oxysporum*, *penicillium chrysogenum* var. *chrysogenum*, *phoma destructiva*, *aureobasidium pullulans* var. *pullutans*, *rhizopus stolonifer* and *pleospora herbarum* injection, solution

TREATMENT SET TS333177 - *poa pratensis* pollen, *ambrosia trifida* pollen, *ambrosia artemisiifolia* pollen, *ambrosia psilostachya* pollen, *cynodon dactylon* pollen, *acer negundo* pollen, *populus deltoides* pollen, *carya illinoiensis* pollen, *carya alba* pollen, *betula nigra* pollen, *fraxinus americana* pollen, *quercus alba* pollen, *sorghum halepense* pollen, *chenopodium album* pollen, *iva annua* var. *annua* pollen, *amaranthus retroflexus* pollen, *salsola kali* pollen and *rumex acetosella* pollen injection, solution

TREATMENT SET TS332486 - *ambrosia trifida* pollen, *ambrosia artemisiifolia* pollen, *ambrosia psilostachya* pollen, *carya illinoiensis* pollen, *carya alba* pollen, *cynodon dactylon* pollen, *plantago lanceolata* pollen, *juniperus virginiana* pollen, *acer negundo* pollen, *populus deltoides* pollen, *betula nigra* pollen, *fraxinus americana* pollen, *quercus alba* pollen, *sorghum halepense* pollen, *kochia scoparia* pollen, *chenopodium album* pollen, *iva annua* var. *annua* pollen, *amaranthus retroflexus* pollen, *salsola kali* pollen and *rumex acetosella* pollen injection, solution

TREATMENT SET TS332487 - *gallus gallus* feather, *anas platyrhynchos* feather, *anser anser* feather, *ustilago nuda hordei*, *ustilago maydis*, *ustilago avenae*, *ustilago tritici*, *ustilago cynodontis*, *sporisorium cruentum*, *cochliobolus sativus*, *penicillium chrysogenum* var. *chrysogenum*, *aureobasidium pullulans* var. *pullutans*, *pleospora herbarum*, *canis lupus familiaris* hair, *periplaneta americana*, *dermatophagoides pteronyssinus*, *candida albicans*, *fusarium oxysporum*, *cladosporium cladosporioides*, *phoma destructiva* and *rhizopus stolonifer* injection, solution

TREATMENT SET TS329227 - *ambrosia trifida* pollen, *ambrosia artemisiifolia* pollen, *ambrosia psilostachya* pollen, *carya illinoiensis* pollen, *carya alba* pollen, *ustilago nuda hordei*, *ustilago maydis*, *ustilago avenae*, *ustilago tritici*, *ustilago cynodontis*, *sporisorium cruentum*, *dermatophagoides pteronyssinus*, *ulmus americana* pollen, *plantago lanceolata* pollen, *cynodon dactylon* pollen, *poa pratensis* pollen, *fraxinus americana* pollen, *betula nigra* pollen, *populus deltoides* pollen, *quercus alba* pollen, *sorghum halepense* pollen, *amaranthus retroflexus* pollen, *salsola kali* pollen, *kochia scoparia* pollen, *iva annua* var. *annua* pollen and *chenopodium album* pollen injection, solution

TREATMENT SET TS330716 - *ambrosia trifida* pollen, *ambrosia artemisiifolia* pollen, *ambrosia psilostachya* pollen, *carya illinoiensis* pollen, *carya alba* pollen, *poa pratensis* pollen, *ulmus americana* pollen, *cynodon dactylon* pollen, *plantago lanceolata* pollen, *juniperus virginiana* pollen, *acer negundo* pollen, *populus deltoides* pollen, *betula nigra* pollen, *fraxinus americana* pollen, *quercus alba* pollen, *sorghum halepense* pollen, *chenopodium album* pollen, *iva annua* var. *annua* pollen and *amaranthus retroflexus* pollen injection, solution

TREATMENT SET TS330717 - *dermatophagoides farinae*, *alternaria alternata*, *canis lupus familiaris* hair, *gallus gallus* feather, *anas platyrhynchos* feather, *anser anser* feather, *dermatophagoides pteronyssinus*, *candida albicans*, *aspergillus fumigatus*, *fusarium oxysporum*, *cochliobolus sativus*, *cladosporium cladosporioides*, *aureobasidium pullulans* var. *pullutans*, *pleospora herbarum*, *rhizopus stolonifer*, *ustilago nuda hordei*, *ustilago maydis*, *ustilago avenae*, *ustilago tritici*, *ustilago cynodontis* and *sporisorium cruentum* injection, solution

TREATMENT SET TS332340 - *gallus gallus* feather, *anas platyrhynchos* feather, *anser anser* feather, *ambrosia trifida* pollen, *ambrosia artemisiifolia* pollen, *ambrosia psilostachya* pollen, *carya illinoiensis* pollen, *carya alba* pollen, *ustilago cynodontis*, *sporisorium cruentum*, *periplaneta americana*, *dermatophagoides farinae*, *dermatophagoides pteronyssinus*, *canis lupus familiaris* hair, *felis catus* hair, *ulmus americana* pollen, *cynodon dactylon* pollen, *sorghum halepense* pollen, *fraxinus americana* pollen, *acer negundo* pollen, *juniperus virginiana* pollen, *poa pratensis* pollen, *rumex acetosella* pollen, *amaranthus retroflexus* pollen, *kochia scoparia* pollen, *iva annua* var. *annua* pollen, *candida albicans*, *cochliobolus sativus*, *rhizopus stolonifer* and *pleospora herbarum* injection, solution

TREATMENT SET TS331258 - *ambrosia trifida* pollen, *ambrosia artemisiifolia* pollen, *ambrosia psilostachya* pollen, *carya illinoiensis* pollen, *carya alba* pollen, *poa pratensis* pollen, *ulmus americana* pollen, *cynodon dactylon* pollen, *plantago lanceolata* pollen, *juniperus virginiana* pollen, *acer negundo* pollen, *populus deltoides* pollen, *betula nigra* pollen, *fraxinus americana* pollen, *quercus alba* pollen, *sorghum halepense* pollen, *kochia scoparia* pollen, *chenopodium album* pollen, *iva annua* var. *annua* pollen, *amaranthus retroflexus* pollen, *salsola kali* pollen and *rumex acetosella* pollen injection, solution

TREATMENT SET TS331259 - *gallus gallus* feather, *anas platyrhynchos* feather, *anser anser* feather, *ustilago cynodontis*, *sporisorium cruentum*, *pleospora herbarum*, *cladosporium cladosporioides*, *dermatophagoides farinae*, *alternaria alternata*, *canis lupus familiaris* hair, *periplaneta americana*, *dermatophagoides pteronyssinus*, *aspergillus fumigatus* and *fusarium oxysporum* injection, solution

TREATMENT SET TS331794 - ustilago cynodontis, sporisorium cruentum, dermatophagoides pteronyssinus, juniperus virginiana pollen, sorghum halepense pollen, chenopodium album pollen, iva annua var. annua pollen, amaranthus retroflexus pollen, salsola kali pollen and rumex acetosella pollen injection, solution

TREATMENT SET TS330533 - gallus gallus feather, anas platyrhynchos feather, anser anser feather, ustilago nuda hordei, ustilago maydis, ustilago avenae, ustilago tritici, ustilago cynodontis, sporisorium cruentum, cochliobolus sativus, cladosporium cladosporioides, pleospora herbarum, aureobasidium pullulans var. pullutans, penicillium chrysogenum var. chrysogenum, felis catus hair, rhizopus stolonifer, dermatophagoides farinae, phoma destructiva, fusarium oxysporum, candida albicans, aspergillus fumigatus, dermatophagoides pteronyssinus, alternaria alternata, canis lupus familiaris hair and periplaneta americana injection, solution

TREATMENT SET TS330534 - carya illinoinensis pollen, carya alba pollen, cynodon dactylon pollen, juniperus virginiana pollen, plantago lanceolata pollen, acer negundo pollen, populus deltoides pollen, iva annua var. annua pollen, chenopodium album pollen, amaranthus retroflexus pollen, betula nigra pollen, fraxinus americana pollen, quercus alba pollen and sorghum halepense pollen injection, solution

TREATMENT SET TS332850 - dermatophagoides farinae, dermatophagoides pteronyssinus, felis catus hair, cladosporium cladosporioides, rumex acetosella pollen and aspergillus fumigatus injection, solution

TREATMENT SET TS332131 - ambrosia trifida pollen, ambrosia artemisiifolia pollen, ambrosia psilostachya pollen, periplaneta americana, gallus gallus feather, anas platyrhynchos feather, anser anser feather, dermatophagoides farinae, dermatophagoides pteronyssinus, canis lupus familiaris hair, felis catus hair, ulmus americana pollen, plantago lanceolata pollen, cynodon dactylon pollen, poa pratensis pollen, alternaria alternata, cladosporium cladosporioides, betula nigra pollen, populus deltoides pollen, amaranthus retroflexus pollen, aureobasidium pullulans var. pullutans, kochia scoparia pollen and candida albicans injection, solution

TREATMENT SET TS332507 - ambrosia trifida pollen, ambrosia artemisiifolia pollen, ambrosia psilostachya pollen, ulmus americana pollen, poa pratensis pollen, alternaria alternata, fraxinus americana pollen, betula nigra pollen, acer negundo pollen, populus deltoides pollen and quercus alba pollen injection, solution

TREATMENT SET TS331567 - ambrosia trifida pollen, ambrosia artemisiifolia pollen, ambrosia psilostachya pollen, carya illinoinensis pollen, carya alba pollen, dermatophagoides farinae, dermatophagoides pteronyssinus, canis lupus familiaris hair, felis catus hair, cynodon dactylon pollen, poa pratensis pollen, alternaria alternata, juniperus virginiana pollen and candida albicans injection, solution

TREATMENT SET TS331133 - ambrosia trifida pollen, ambrosia artemisiifolia pollen, ambrosia psilostachya pollen, carya illinoinensis pollen, carya alba pollen, ustilago nuda hordei, ustilago maydis, ustilago avenae, pleospora herbarum, ustilago cynodontis, sporisorium cruentum, felis catus hair, poa pratensis pollen, alternaria alternata, cynodon dactylon pollen, dermatophagoides pteronyssinus, juniperus virginiana pollen, acer negundo pollen, populus deltoides pollen, betula nigra pollen, fraxinus americana pollen, quercus alba pollen, sorghum halepense pollen, kochia scoparia pollen, chenopodium album pollen, amaranthus retroflexus pollen and salsola kali pollen injection, solution

TREATMENT SET TS333374 - ambrosia trifida pollen, ambrosia artemisiifolia pollen, ambrosia psilostachya pollen, poa pratensis pollen, cynodon dactylon pollen, plantago lanceolata pollen, juniperus virginiana pollen, acer negundo pollen, betula nigra pollen, fraxinus americana pollen, sorghum halepense pollen, iva annua var. annua pollen and chenopodium album pollen injection, solution

TREATMENT SET TS333373 - gallus gallus feather, anas platyrhynchos feather, anser anser feather, ustilago nuda hordei, ustilago maydis, ustilago avenae, ustilago tritici, cladosporium cladosporioides, cochliobolus sativus, aureobasidium pullulans var. pullutans, penicillium chrysogenum var. chrysogenum, pleospora herbarum, felis catus hair, dermatophagoides farinae, alternaria alternata, canis lupus familiaris hair, periplaneta americana, dermatophagoides pteronyssinus, candida albicans, aspergillus fumigatus, fusarium oxysporum, phoma destructiva and rhizopus stolonifer injection, solution

TREATMENT SET TS332287 - carya illinoinensis pollen, carya alba pollen, ambrosia trifida pollen, ambrosia artemisiifolia pollen, ambrosia psilostachya pollen, poa pratensis pollen, ulmus americana pollen, cynodon dactylon pollen, plantago lanceolata pollen, acer negundo pollen, populus deltoides pollen, betula nigra pollen, fraxinus americana pollen, quercus alba pollen, sorghum halepense pollen and iva annua var. annua pollen injection, solution

TREATMENT SET TS332291 - gallus gallus feather, anas platyrhynchos feather, anser anser feather, ustilago nuda hordei, ustilago maydis, ustilago avenae, ustilago tritici, pleospora herbarum, aureobasidium pullulans var. pullutans, cladosporium cladosporioides, cochliobolus sativus, rhizopus stolonifer, phoma destructiva, fusarium oxysporum, aspergillus fumigatus, dermatophagoides farinae, dermatophagoides pteronyssinus, alternaria alternata, canis lupus familiaris hair and periplaneta americana injection, solution

TREATMENT SET TS329240 - ambrosia trifida pollen, ambrosia artemisiifolia pollen, ambrosia psilostachya pollen, carya illinoinensis pollen, carya alba pollen, pleospora herbarum, penicillium chrysogenum var. chrysogenum, juniperus virginiana pollen, quercus alba pollen, fraxinus americana pollen, ulmus americana pollen, poa pratensis pollen, cynodon dactylon pollen, dermatophagoides pteronyssinus, plantago lanceolata pollen, phoma destructiva, candida albicans, kochia scoparia pollen, acer negundo pollen, populus deltoides pollen, betula nigra pollen, sorghum halepense pollen, chenopodium album pollen, iva annua var. annua pollen, amaranthus retroflexus pollen and salsola kali pollen injection, solution

TREATMENT SET TS333375 - ambrosia psilostachya pollen, ambrosia artemisiifolia pollen, ambrosia trifida pollen, carya alba pollen, carya illinoinensis pollen, juniperus virginiana pollen, quercus alba pollen, ulmus americana pollen, plantago

lanceolata pollen, cynodon dactylon pollen, poa pratensis pollen, fraxinus americana pollen, betula nigra pollen, acer negundo pollen, populus deltoides pollen, sorghum halepense pollen, rumex acetosella pollen, amaranthus retroflexus pollen, salsola kali pollen, kochia scoparia pollen, iva annua var. annua pollen and chenopodium album pollen injection, solution

TREATMENT SET TS330821 - ambrosia trifida pollen, ambrosia artemisiifolia pollen, ambrosia psilostachya pollen, ustilago cynodontis, sporisorium cruentum, gallus gallus feather, anas platyrhynchos feather, anser anser feather, juniperus virginiana pollen, ulmus americana pollen, plantago lanceolata pollen, felis catus hair, poa pratensis pollen, dermatophagoides farinae, alternaria alternata, periplaneta americana, cynodon dactylon pollen, dermatophagoides pteronyssinus, candida albicans, acer negundo pollen, betula nigra pollen, sorghum halepense pollen, chenopodium album pollen and iva annua var. annua pollen injection, solution

TREATMENT SET TS330156 - ambrosia psilostachya pollen, ambrosia artemisiifolia pollen, ambrosia trifida pollen, carya illinoinensis pollen, carya alba pollen, sporisorium cruentum, ustilago cynodontis, ustilago nuda hordei, ustilago maydis, ustilago avenae, ustilago tritici, pleospora herbarum, cladosporium cladosporioides, quercus alba pollen, fraxinus americana pollen, ulmus americana pollen, felis catus hair, poa pratensis pollen, alternaria alternata, cynodon dactylon pollen, candida albicans, fusarium oxysporum, acer negundo pollen, betula nigra pollen, sorghum halepense pollen, kochia scoparia pollen and salsola kali pollen injection, solution

TREATMENT SET TS332678 - ustilago cynodontis, sporisorium cruentum, gallus gallus feather, anas platyrhynchos feather, anser anser feather, penicillium chrysogenum var. chrysogenum, cladosporium cladosporioides, felis catus hair, dermatophagoides farinae, alternaria alternata, dermatophagoides pteronyssinus and aspergillus fumigatus injection, solution

TREATMENT SET TS332679 - ambrosia trifida pollen, ambrosia artemisiifolia pollen, ambrosia psilostachya pollen, carya alba pollen, carya illinoinensis pollen, juniperus virginiana pollen, quercus alba pollen, fraxinus americana pollen, ulmus americana pollen, plantago lanceolata pollen, poa pratensis pollen, cynodon dactylon pollen, acer negundo pollen, rumex acetosella pollen, chenopodium album pollen, sorghum halepense pollen, betula nigra pollen, populus deltoides pollen, amaranthus retroflexus pollen, salsola kali pollen, kochia scoparia pollen and iva annua var. annua pollen injection, solution

TREATMENT SET TS331795 - sporisorium cruentum, ustilago cynodontis, ustilago nuda hordei, ustilago maydis, ustilago avenae, ustilago tritici, gallus gallus feather, anas platyrhynchos feather, anser anser feather, pleospora herbarum, penicillium chrysogenum var. chrysogenum, cochliobolus sativus, aureobasidium pullulans var. pullutans, cladosporium cladosporioides, rhizopus stolonifer, dermatophagoides farinae, dermatophagoides pteronyssinus, phoma destructiva, periplaneta americana, candida albicans, aspergillus fumigatus and fusarium oxysporum injection, solution

TREATMENT SET TS331796 - carya illinoinensis pollen, carya alba pollen, juniperus virginiana pollen, quercus alba pollen, fraxinus americana pollen, cynodon dactylon pollen, acer negundo pollen, populus deltoides pollen, betula nigra pollen, chenopodium album pollen, iva annua var. annua pollen, amaranthus retroflexus pollen, salsola kali pollen and rumex acetosella pollen injection, solution

TREATMENT SET TS329413 - cochliobolus sativus, fraxinus americana pollen, periplaneta americana, dermatophagoides farinae, dermatophagoides pteronyssinus, salsola kali pollen, iva annua var. annua pollen and candida albicans injection, solution

TREATMENT SET TS334696 - rumex acetosella pollen, urtica dioica pollen, felis catus hair, canis lupus familiaris hair, acer negundo pollen, juniperus ashei pollen, juglans nigra pollen, platanus occidentalis pollen, populus deltoides pollen, fraxinus americana pollen, quercus alba pollen, ulmus americana pollen, morus alba pollen, cynodon dactylon pollen, phleum pratense pollen, ambrosia artemisiifolia pollen, amaranthus retroflexus pollen, salsola kali pollen, iva annua var. annua pollen, carya alba pollen and carya illinoinensis pollen injection, solution

Antigen Laboratories, Inc.

## WARNINGS

Individual allergenic extract treatment vial is intended for use by physicians who are experienced in the administration of allergenic extracts for immunotherapy and the emergency care of anaphylaxis, or for use under the guidance of an allergy specialist. Individual allergenic extract treatment vials are not directly interchangeable with other allergenic extracts. The initial dose must be based on skin testing as described in the dosage and administration section of this insert. Patients being switched from other types of extract to individual treatment vials should be started as though they were coming under treatment for the first time. Patients should be instructed to recognize adverse reaction symptoms and cautioned to contact the physician's office if reaction symptoms occur. As with all allergenic extracts, severe systemic reactions may occur. In certain individuals, these reactions may be life-threatening. Patient should be observed for at least 20 minutes following treatment and emergency measures as well as personnel trained in their use should be immediately available in the event of a life-threatening reaction.

This product should not be injected intravenously. Deep subcutaneous routes have proven to be safe. See the warnings, precautions, adverse reactions and overdosage sections below.

## **DESCRIPTION**

Allergenic extract in this vial is referred to as an individual treatment vial since it is designed primarily for the physician equipped to complete skin testing and supervise allergenic extract immunotherapy. The extract is sterile and intended for subcutaneous injection. The concentration of allergenic extract supplied will be based on the individual physician's prescription order and will be expressed in most cases on a weight/volume basis (or AU/ml with standardized extract) diluted either 1:10 or 1:5. Where mixtures of pollens and non-pollens have been ordered, the ingredients are listed on the final container label. To insure maximum potency for the entire dating period, dilutions will be prepared with 50% v/v glycerine unless otherwise specified.

**Ingredients** - Active allergens, preservative and stabilizer are noted on the Physicians Prescription Ingredients Insert enclosed with each individual allergenic extract treatment vial.

**Dating Period** - A twelve month dating period (expiration date) for the prescription vial will be on the container label. Extract Treatment Sets should be reordered when outdated. Government requirements include a two week holding period for sterility tests. Please allow three weeks minimum for delivery.

## **CLINICAL PHARMACOLOGY**

The mechanisms by which immunotherapy (hypersensitization) is achieved are not completely understood. Anaphylaxis or "anaphylactic shock," and hay fever are caused by the same basic process: the production of IgE antibody, its attachment to mast cells and, on renewed contact with the same antigen explosive degranulation of the mast cells and release of mediators, which act on smooth muscle, mucous glands, and blood vessels. With massive release there is bronchospasm, vomiting, skin rashes, edema of the nose and throat, and vascular collapse, sometimes fatal, while with more localized release one or more of these symptoms predominates, depending on the site (tissue shock organ) of exposure to the antigen.

Antigens that can trigger these reactions are known as "allergens"; they have very diverse origins but a curious similarity of molecular weight. People who suffer unduly from allergy are called "atopic"; this trait is usually inherited and has been attributed to a variety of constitutional abnormalities.

The IgE dependent degranulation of mast cells is initiated by the bridging of pairs of cell-bound IgE by antigen and terminates rapidly. Bridging results in alteration of the cell membrane, which is associated with increased energy dependent entry of calcium, alterations in phospholipid metabolism and increase of cyclic adenosine monophosphate (cyclic AMP).

The mast cell membrane is ruffled and possesses receptors both for the Fc portion of IgE and C3b. Receptors for anaphylatoxin (C3a and C5a), have been defined functionally. In addition to IgE antigen interaction and stimulation by anaphylatoxin, mast cells may be degranulated by non-immunological stimuli such as enzymes, ionophores, polycations, radio-contrast dyes and opiates. Atopic individuals develop their symptoms principally as a result of IgE-dependent processes; however, non- IgE mediated mechanisms for the release of mast cell mediators provide additional potential for recruitment of mediators.

Subsequent to activation, the secretion of granules is under cyclic nucleotide regulation. Of direct relevance is the possibility that the mast cell itself, by histamine ( $H^2$ ) and prostaglandins ( $E^2, D^2, I^2$ ) may increase cyclic AMP and inhibit secretion. Conversely histamine ( $H^1$ ) could elevate cyclic GMP and PGF, 2 alpha lower cyclic AMP, augmenting the release of mediators. 18

## **INDICATIONS AND USAGE**

When the natural exposure to elevated aeroallergens produces symptoms as described under Clinical Pharmacology, specific diagnosis and therapeutic procedures are indicated. Clearly, important clues to the cause of a person's allergic condition can be gleaned from a thorough history and careful physical examination. Diagnostic tests - in vitro or in vivo - serve only to confirm the physician's suspicions or to improve investigative skills. Specific diagnosis is especially indicated when the patient's symptoms are not controlled by medication. When immunotherapy is contemplated demonstration of sensitivity to a specific allergenic extract is necessary. An orderly approach to the use of diagnostic tests usually begins with direct skin testing. 5,6,11

**THIS PRODUCT IS NOT INTENDED FOR TREATMENT OF PATIENTS WHO DO NOT MANIFEST IMMEDIATE HYPERSENSITIVITY REACTIONS TO THE ALLERGENIC EXTRACT FOLLOWING SKIN TESTING.**

## **CONTRAINdicATIONS**

There are no absolute contraindications; however, extreme caution is necessary when using diagnostic skin tests or injection treatment in highly sensitive patients, who have experienced severe symptoms or anaphylaxis by natural exposure or previous skin testing or treatment. **IN THESE CASES BOTH THE POTENCY FOR SKIN TESTS AND THE ESCALATION OF THE TREATMENT DOSE MUST BE ADJUSTED TO THE PATIENT'S SENSITIVITY AND TOLERANCE.**

This product is not intended for the treatment of patients who do not experience allergic symptoms upon natural exposure to the allergen. At the present time there has been no demonstrated adverse effects on the fetus when allergenic extract immunotherapy is administered during gestation to pregnant women.

**100,000 AU/ml standardized allergenic extract should be used by physicians with experience in maximal dose immunotherapy and treatment of anaphylaxis.**

## **WARNINGS**

Epinephrine 1:1000 should be available.

When changing immunotherapy from an unstandardized to an AU/ml standardized allergenic extract, dose adjustment, if indicated, should be based on the comparative potency of the extracts. Patient re-evaluation may be necessary.

Injections should never be given intravenously. A 5/8 inch 25 gauge needle on a sterile syringe will allow deep subcutaneous injection. Precaution of withdrawing the plunger slightly after inserting the needle is advisable to determine if a blood vessel has been entered. Proper measurement of the dose and caution in making the injection will minimize reactions. Patients should be detained for twenty to thirty minutes after injection or advised to return to the office immediately if symptoms or reactions occur.

**Sensitive patients may experience severe anaphylactic reactions resulting in respiratory obstruction, shock, coma and/or death.**

## GENERAL PRECAUTIONS

Pregnancy Category C: There are no adequate and well controlled studies in pregnant women. Allergenic extracts should be used during pregnancy only if the potential benefit justifies the potential risk to the fetus.

Pediatric Use: Doses of allergenic extracts for children are generally the same as those for adults. The maximum volume of extract tolerated without undue pain and swelling may be less for smaller individuals and therefore the maximum dose and treatment schedule must be individualized.

Animal reproduction studies have not been conducted with allergenic extracts. It is also not known whether allergenic extracts can cause fetal harm when administered to a pregnant woman or can affect reproduction capacity. Allergenic extracts should be given to a pregnant woman only if clearly needed.<sup>15</sup>

Refrigerate at 2-8 degrees C.

Sterile solutions, vials, syringes, etc. should be used. Aseptic technique should be observed when making dilutions, skin testing, and extracts for treatment. The usual precautions in administering allergenic extracts are necessary.

A separate autoclave sterilized or disposable syringe and needle should be used for each individual patient to prevent transmission of serum hepatitis, A.I.D.S. and other infectious agents from one person to another.

The initial dilution of allergen extract, starting dose, and progression of dosage must be carefully determined on the basis of the patient's history and results of skin tests. Patients with a history of severe sensitivity and markedly positive skin tests to high dilutions of the allergen extract should be started with low doses of highly diluted extract. Pregnancy or a history of prior reactions to allergen immunotherapy dictates the need to start with small quantities of antigen.

If the first injection of the initial dilution of extract is tolerated without significant local reaction, increasing doses by 5-20% increments of that dilution may be administered. The rate of increase in dosage in the early stages of treatment with highly diluted extracts is usually more rapid than the rate of increase possible with more concentrated extracts. This schedule is intended only as a guide and must be modified according to the reactivity of the individual patient. Needless to say, the physician must proceed cautiously in the treatment of the highly sensitive patient who develops large local or systemic reactions.

Some patients may tolerate larger doses, but it is rarely necessary to give maintenance doses larger than 2,000 AU/ml of the standardized extract or 0.5 ml of 1:100 w/v of the unstandardized extract. Because dilute extracts tend to lose activity on storage, the first dose from a more concentrated vial, should be the same or less than the previous dose.<sup>1,2,3,4,16</sup>

Immunotherapy must be given under the supervision of a physician. Before an injection is given the patient should be asked about any reaction following the previous injection to help determine the next dose. Target maintenance dose should be determined by the physician's experience and patient response to skin testing and treatment.

Dosages progressively increase thereafter according to the tolerance of the patient at intervals of one to seven days until, (1) the patient achieves relief of symptoms, (2) induration at the site of injection is no larger than 50 mm in 36 to 48 hours, (3) a maintenance dose short of aggravating existing symptoms, systemic symptoms, or anaphylaxis. The dose should be escalated until the patient is receiving a final maintenance dose containing 2.7 to 22 u/ml or more of ragweed AgE (using Short Ragweed as example) or demonstrates untoward reactions that indicate the dose to be excessive. This maintenance dose may be continued at regular intervals perennially or achieved each year by a new, but shortened course of treatment. It may be necessary to adjust the progression of dosage downward to avoid local and constitutional reactions.

Immunotherapy of patients highly sensitive to ragweed pollen (using Short Ragweed as an example) receiving a dose of 2.7 to 46.8 units of ragweed AgE (1,000-2,000 or more AU/ml of standardized ragweed extract) was significantly more effective than placebo for (1) relieving symptoms of ragweed hay fever, (2) producing increase in serum levels of anti-ragweed IgG, (3) decrease in seasonal rises in levels of anti-ragweed IgE, (4) decrease in leukocyte histamine release from exposure to ragweed pollen extract in some patients, and (5) increase in IgG and IgA antibodies in nasal secretions.<sup>17</sup>

In addition to these changes in humoral antibody production, immunotherapy also effects some cellular changes. Basophils from treated subjects release less histamine in vitro and are less sensitive to the allergen (that is, higher concentrations of allergens are required to induce histamine release) than are basophils from non-treated patients. Lymphocytes from treated patients exhibit decreased proliferative response and decreased production of lymphokine in the presence of the specific allergen. A state of tolerance may be induced in the IgE producing B lymphocytes, there may be impairment in T-lymphocyte helper function, or immunotherapy may generate suppressor cells. Antigen specific suppressor cells, probably bearing histamine receptors, are generated during immunotherapy for allergy and may be partly responsible for the efficacy of this therapy.<sup>14</sup>

Loss of potency of aqueous pollen extracts has been recognized as a problem since shortly after the introduction of modern methods of immunotherapy. This loss of potency occurs more rapidly in saline extracts without added preservatives at high temperature and

at greater extract dilutions. At concentrations of 1:100 all dilutions containing glycerin, human serum albumin, maintained extract potency within 1 logarithm (log) dilution of the original strength for 12 months; glycerin was significantly superior to all other extracts at 1, 3, and 12 months; and the deleterious effect of phenol was minimal. The deleterious effect of phenol was more marked at the higher dilutions. It was concluded that there may be marked loss of potency of dilute pollen extracts stored for periods of only two weeks under conditions which may be encountered in normal clinical practice. 12

## DOSAGE AND ADMINISTRATION

As a consequence of the discovery of IgE and the development of methods to identify and quantify anti-allergen IgE levels, interest in recent years has centered around the utilization of in vivo and in vitro diagnostic procedures. 7,9

Patients who react to a small quantity of antigen by skin testing can be classified as highly sensitive. Those who react only to large quantities of antigen can be classified as less sensitive. It would appear that there is at least a 50,000-fold range between the most and least sensitive individuals. On the other hand, certain patients who do not appear to have elevated quantities of specific anti-allergen IgE do have positive skin tests and have symptoms of allergic rhinitis. These patients are considerably less sensitive than patients with detectable levels of specific IgE antibody. 10

The current standard method of immunotherapy dates back to the earliest studies by Noon. As adapted for ragweed pollen extract, therapy is begun with a low dose, which has been shown to be tolerated by both experience and skin testing.

The physician who undertakes immunotherapy should be concerned with the degree of sensitivity of the patient. This can be measured by skin test, leukocyte histamine release, or anti-allergen IgE levels. Strongly positive skin tests or high initial ragweed IgE and total IgE may be risk factors for systemic reactions. Less aggressive immunotherapy schedules may be indicated for such patients. Maintenance dose potency must be established by the physician's clinical observation and experience. 10,17

Serial fivefold or tenfold dilutions of the extract are used to make more dilute extract concentrations. Other concentrations can be prepared by appropriate dilution. In brief, the allergist can prepare any dilution of extract that is considered appropriate for the patient.

## ADVERSE REACTIONS

Systemic reactions may range from mild exaggeration of the patient's allergic symptoms to anaphylactic reactions. Very sensitive patients may show a rapid response. In some instances a severe systemic reaction with blood pressure fall and/or shock may occur. Quantitation of patient's sensitivity combined with careful early observation is essential for safe skin testing and treatment.

Other reactions include, but are not necessarily limited to urticaria, itching, edema of the extremities, respiratory wheezing or asthma, dyspnea, cyanosis, tachycardia, lacrimation, marked perspiration, flushing of the face, neck or upper chest, mild persistent clearing of the throat, hacking cough, or persistent sneezing.

### 1) Local Reactions

Small amounts of erythema and swelling at the site of injection are common, the extent varying with the patient. Such reactions should not be considered significant unless they persist for at least 24 hours or exceed 50 mm. in diameter.

Larger local reactions are not only uncomfortable, but also indicate the possibility of a systemic reaction if dosage is increased. In such cases the dosage should be reduced to the last level not causing the reaction and maintained at this level for two or three treatments before cautiously increasing again.

Large, persistent local reactions or minor exacerbations of the patient's allergic symptoms may be treated by local cold applications and/or the use of oral antihistamines, but they should be considered a warning of possible severe systemic reactions and the need for temporarily reduced dosages.

A mild burning immediately after the injection is to be expected; this usually leaves in 10 to 20 seconds. Prolonged pain, or pain radiating up the arm, is usually the result of intramuscular injection, making this injection route undesirable. Subcutaneous injection is the recommended route.

### 2) Systemic Reactions

With careful attention to dosage and administration, such reactions occur infrequently, but it must be remembered that allergenic extracts are highly potent to sensitive individuals and OVERDOSE could result in anaphylactic symptoms. Therefore, it is imperative that physicians administering allergenic extracts understand and be prepared for the treatment of severe reactions.

Adverse reaction frequency data for allergenic extract administration is not available. Inherent difficulties in establishing such data are the wide variations in clinical allergy types, patient sensitivity, treatment schedules used by allergists, potency of extracts from various sources, etc.

It cannot be overemphasized that, under certain unpredictable combinations of circumstances, anaphylactic shock is always a possibility. Other possible systemic reaction symptoms are, in varying degrees of severity, fainting, pallor, bradycardia, hypotension, angioedema, cough, wheezing, conjunctivitis, rhinitis, and urticaria. 17,18

## OVERDOSAGE

If a systemic or anaphylactic reaction does occur, apply a tourniquet above the site of injection and inject intramuscularly or subcutaneously 0.3 to 0.5 ml of 1:1000 epinephrine-hydrochloride into the opposite arm. The dose may be repeated in 5-10 minutes if necessary. Loosen the tourniquet at least every 10 minutes.

The epinephrine HCL 1: 1000 dose for infants to 2 years is 0.05 to 0.1 ml; for children 2 to 6 years it is 0.15 ml; for children 6 to 12 years it is 0.2 ml.

Patients unresponsive to epinephrine may be treated with theophylline. Studies on asthmatic subjects reveal that plasma concentrations of theophylline of 5 to 20 ug/ml are associated with therapeutic effects. Toxicity is particularly apparent at concentrations greater than 20 ug/ml. A loading dose of aminophylline of 5.6 mg/kg intravenously followed by 0.9 mg/kg per hour results in plasma concentrations of approximately 10 ug/ml. (Mitenko and Ogilive 1973b; Nicholoson and Chick 1973).

Other beta-adrenergic drugs such as isoproterenol, isoetharine, or albuterol may be used by inhalation. The usual dose to relieve broncho-constriction in asthma is 0.5 ml or the 0.5% solution for isoproterenol HCL; albuterol is longer acting than isoproterenol by any route of administration. The albuterol inhaler delivers approximately 90 mcg of albuterol from the mouthpiece. The usual dosage for adults and children would be two inhalations repeated every 4 to 6 hours. Isoetharine supplied in the Bronkometer unit delivers approximately 340 mcg isoetharine. The average adult dose is one to two inhalations.

**Patients receiving beta-blockers may not be responsive to epinephrine or inhaled bronchodilators.** Respiratory obstruction not responding to parenteral or inhaled bronchodilators may require Theophylline, low-flow oxygen, intubation and the use of life support systems. Parenteral fluid and/or plasma expanders may be utilized for treatment of shock and low flow (two liters per minute) oxygen may be utilized if indicated. Adrenocorticosteroids may be administered parenterally or intravenously. 8

## HOW SUPPLIED

Individual treatment sets as prescribed by the physician. The allergenic extract contains a variable number of individual doses depending on the patient's sensitivity and maximum tolerated maintenance treatment dose.

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CONTAINER LABELING

# ALLERGENIC EXTRACT

Initial dose: 0.10 ml **subcutaneous**.  
CAUTION: U.S. Federal Law prohibits dispensing without prescription.

U.S. Government License No. 468  
No U.S. Standard of Potency  
**REFRIGERATE AT 2°-8° C.**  
**NON-RETURNABLE**

For Physician's Use Only. See Physician's Rx ingredients insert for active allergens, preservative and stabilizer.

**WARNING:** Read Individual Treatment Vial insert for instructions, dilution and dosage schedule.  
P.O. BOX 123, LIBERTY, MO 64069 U.S.A.



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CAUTION: U.S. Federal Law prohibits dispensing without prescription.

U.S. Government License No. 468  
No U.S. Standard of Potency  
**REFRIGERATE AT 2°-8° C.**  
**NON-RETURNABLE**

For Physician's Use Only. See Physician's Rx ingredients insert for active allergens, preservative and stabilizer.  
**WARNING:** Read Individual Treatment Vial insert for instructions, dilution and dosage schedule.  
P.O. BOX 123, LIBERTY, MO 64069 U.S.A.

